## **Deconvolution Approach to Modeling**

#### Deconvolution + regression

- ★ What is deconvolution in the context of FMRI regression?
- ★ How different from shape-prefixed approach?
- ⋆ Pros and cons
- ★ Implementation via TENT basis function in 3dDeconvolve
- ⋆ Collinearity issue
- ★ Complications for group analysis

#### Example of HRF modeling with TENT basis function

- ⋆ Preprocessing steps
- ★ Regression model with 3dDeconvolve
- ★ Stimulus timing setup
- ★ Specifications for general linear testing: *t* and *F*

#### Deconvolution vs. Convolution

#### Convolution + regression

- ★ Impulse response © stimulus = BOLD response (speed X time = dist.)
- ★ Known: impulse response, stimulus, and BOLD response to a stimulus
- ★ Use convolution to create regressors (waver or 3dDeconvolve)
- $\star$  Then response strength ( $\beta$ ) estimated via regression
- ★ Regression with each condition modeled with one basis function

#### <u>Deconvolution</u> + regression

- ★ Deconvolution: derive impulse response or stimulus when the other 2 known in impulse response © stimulus = BOLD response
- ★ Known: stimulus and BOLD response; unknown: impulse response
- ★ Have to estimate average BOLD response to stimuli
- ★ Set a yardstick at each sliding window
- ★ Stimulus response as a linear combination of multiple yardsticks
- ★ Combination of multiple yardsticks © stimulus = regressors
- $\star$  Deconvolution: combination weights ( $\beta$ ) estimated via regression

## Fixed vs. flexible shape modeling

- Basic or Fixed-shape regression (previous talks):
  - ★ Conservative: fix the shape of the HRF amplitude varies
  - ★ Generally used in block or some event-related experiments
  - ★ Use -stim times to generate regressors from stimulus timing
  - $\star$  Find "average" response amplitude in each voxel solution to the set of linear equations =  $\beta$  weights

#### Deconvolution or Variable-shape regression (now):

- ★ Liberal: Allow HRF shape to vary in each voxel, for each stimulus class
- ★ Generally used in event-related or occasionally some block designs
- ★ Appropriate when you don't want to over-constrain the solution by assuming an HRF shape
- $\star$  *Caveat*: need to have enough time points during the HRF in order to resolve its shape (e.g., TR ≤ 3 s)

#### There's always a middle ground

★ Moderate: some wiggle room in shape modeling

### Deconvolution: Pros & Cons (+ & -)

- + Likely to have more accurate response estimate: allowing for subject and regional variability in hemodynamics
- + Can test subtle shape difference (e.g., later time points more "active" than earlier? Undershoot?)
- Need to estimate more parameters for each stimulus class (e.g., 4-15 vs. 1)
- Need more data to get same statistical power (assuming fixed-shape model used was in fact "correct")
- May lead to weird HRF shapes that are difficult to interpret
- Overfitting: noise, head motion, or other regressors may leak into shape estimate
- Challenging for group analysis with multiple βs per regressor: 3dLME

#### Expressing HRF via Regression Unknowns

 The tool for expressing an unknown function as a finite set of numbers that can be fit via linear regression is an <u>expansion in basis functions</u>

$$h(t) = \beta_0 \psi_0(t) + \beta_1 \psi_1(t) + \beta_2 \psi_2(t) + \dots = \sum_{q=0}^{q-p} \beta_q \psi_q(t)$$

- $\star$  Basis functions  $\psi_q(t)$  & expansion order p are known
  - o Larger  $p \Rightarrow$  more complex shapes & more parameters
- $\star$  Unknowns: weight  $\beta_q$  for each  $\psi_q(t)$
- Regressors created through convolution of stimulus timing with h(t)
- βs solved with the regressors in 3dDeconvolve

#### 3dDeconvolve with "Tent Functions"

- Need to describe HRF shape and magnitude with a finite number of parameters
  - ★ Convolution: allows for calculation of h(t) at any arbitrary point in time after the stimulus times:

$$r_n = \sum_{k=1}^{K} h(t_n - \tau_k) = \text{sum of HRF copies}$$

- Simplest set of such functions are <u>tent functions</u>
  - ★ Also known as "piecewise linear splines"

$$T(x) = \begin{cases} 1 - |x| & \text{for } -1 < x < 1 \\ 0 & \text{for } |x| > 1 \end{cases}$$

$$N.B.: \text{ cubic splines are also available}$$

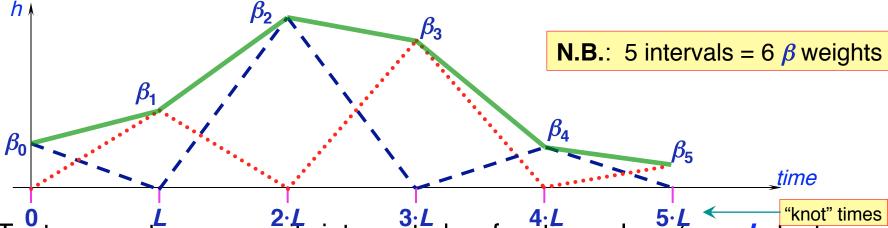
$$t = 0 \qquad t = TR \qquad t = 2 \cdot TR \qquad t = 3 \cdot TR \qquad t = 4 \cdot TR \qquad t = 5 \cdot TR \qquad t = 5 \cdot TR$$

### A

## Tent Functions = Linear Interpolation

 Expansion of HRF in a set of spaced-apart tent functions is the same as linear interpolation between "knots"

$$h(t) = \beta_0 \cdot T\left(\frac{t}{L}\right) + \beta_1 \cdot T\left(\frac{t-L}{L}\right) + \beta_2 \cdot T\left(\frac{t-2\cdot L}{L}\right) + \beta_3 \cdot T\left(\frac{t-3\cdot L}{L}\right) + \cdots$$



- Tent parameters are easily interpreted as function values (e.g., L: tent radius;  $\beta_2$  = response (tent height) at time  $t = 2 \cdot L$  after stim = tent width)
- User must decide on relationship of tent function grid spacing L and time grid spacing TR (usually would choose L ≥ TR)
- In 3dDeconvolve: specify duration (D) of HRF and number (n) of  $\beta$  parameters (details shown a few slides ahead): radius L = D/(n-1) in (n-2) full tents + 2 half tents, each tent overlaps half with two neighboring ones

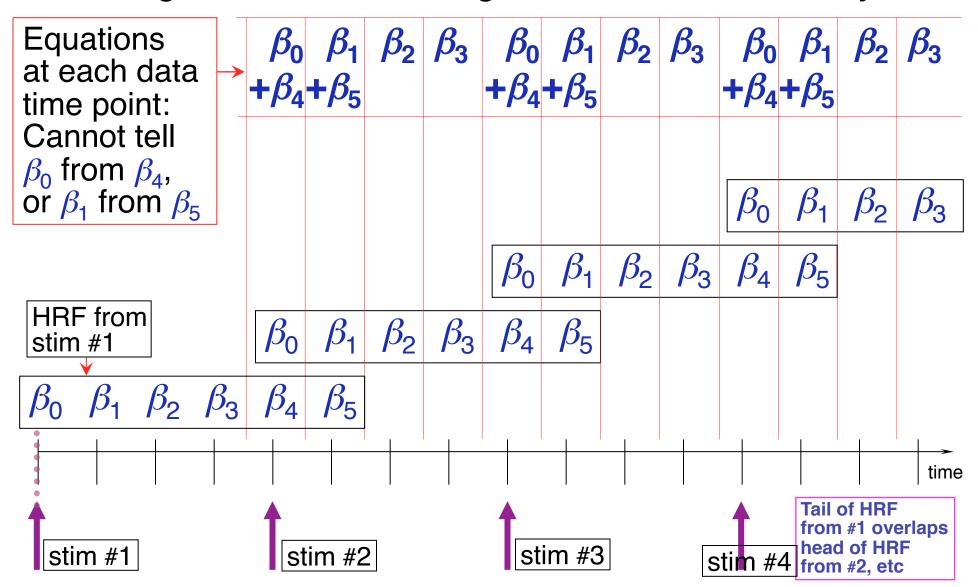
## Tent Functions: Average Signal Change

- For group analysis, may compute average signal change
  - \* Over entire duration of HRF (usual): in previous slide, with 6  $\beta$  weights, average signal change is  $\frac{1}{2}\beta_0 + \beta_1 + \beta_2 + \beta_3 + \beta_4 + \frac{1}{2}\beta_5$
  - ★ Over a sub-interval of the HRF duration (sometimes)
- Or feed all βs into group analysis using 3dLME so that serial correlation is accounted for
  - ⋆ More cumbersome than fixed-shape modeling
- First and last β weights are scaled by half since they only affect half as much of the duration of the response
  - ★ To assume a HRF of D seconds duration with 0 value at beginning & end, use TENT(L, D-L, n-2), not TENT(0, D, n)
- All  $\beta$  weights (for each stimulus class) are output into the "bucket" dataset produced by 3dDeconvolve

## A

## **Deconvolution and Collinearity**

Regular stimulus timing can lead to collinearity!



#### <u>Deconvolution Example - The Data</u>

- cd AFNI\_data2
  - ★ data is in ED/ subdirectory (10 runs of 136 images each; TR=2 s)
  - \* SCript = s1.afni\_proc\_command (in AFNI\_data2/ directory)
    - o stimuli timing and GLT contrast files in misc\_files/
  - \* this script runs program afni\_proc.py to generate a shell script with all AFNI commands for single-subject analysis
    - Run by typing tcsh s1.afni\_proc\_command; then copy/paste
       tcsh -x proc.ED.8.glt | & tee output.proc.ED.8.glt

Text output from

screen and file

programs goes to

- Event-related study from Mike Beauchamp
  - ★ 10 runs, four classes (2x2 design) of stimuli (short videos)
    - Tools moving (e.g., a hammer pounding) <u>ToolMovie</u>
    - People moving (e.g., jumping jacks) <u>HumanMovie</u>
    - Points outlining tools moving (no objects, just points) <u>ToolPoint</u>
    - Points outlining people moving <u>HumanPoint</u>
  - ★ Goal: find brain area that distinguishes natural motions (HumanMovie and HumanPoint) from simpler rigid motions (ToolMovie and ToolPoint)

#### Master Script for Data Analysis

```
√ Master script program

afni proc.py
 -dsets ED/ED r??+orig.HEAD

√ 10 input datasets

 -subj id ED.8.glt
                                                       \ Copy anat to output dir
 -copy anat ED/EDspgr
                                                       \ → Discard first 2 TRs
 -tcat remove first trs 2
 -volreg align to first

√ Where to align all EPIs

                                                       \ ✓ Stimulus timing files (4)
 -regress stim times misc files/stim times.*.1D

√ Stimulus labels

 -regress stim labels ToolMovie HumanMovie
                       ToolPoint HumanPoint
                                                        \ HRF model
 -regress basis 'TENT(0,14,8)' Or 'TENT(2,14,6)'
                                                       \ Specifies that next
 -regress opts 3dD
                                                           lines are options to be
 -gltsym ../misc files/glt1.txt -glt label 1 FullF
                                                           passed to
 -gltsym ../misc files/glt2.txt -glt label 2 HvsT
                                                           3dDeconvolve
 -gltsym ../misc files/glt3.txt -glt label 3 MvsP
                                                           directly (in this case,
 -gltsym ../misc files/glt4.txt -glt label 4 HMvsHP
                                                           the GLTs we want
                                                           computed)
 -gltsym ../misc files/glt5.txt -glt label 5 TMvsTP \
 -gltsym ../misc files/glt6.txt -glt label 6 HPvsTP \
 -gltsym ../misc files/glt7.txt -glt label 7 HMvsTM
```

This script generates file proc. ED.8.glt (180 lines), which contains all the AFNI commands to produce analysis results into directory ED.8.glt.results/ (148 files)

## Shell Script for Deconvolution - Outline

- Copy datasets into output directory for processing
- Examine each imaging run for outliers: 3dToutcount
- Time shift each run's slices to a common origin: 3dTshift
- Registration of each imaging run: 3dvolreg
- Bluring: 3dmerge, 3dBlurToFWHM, 3dBlurInMask
- Create a brain mask: 3dAutomask and 3dcalc
- Rescale each voxel time series in each imaging run so that its average through time is 100: 3dTstat and 3dcalc
  - $\star \beta_q$  of 1.6 indicates 1.6% change at tent function knot #q after stimulus
  - \* Biophysics: believe % signal change is relevant physiological parameter
- Catenate all runs into 1 dataset (1360 time points): 3dTcat
  - ★ This dataset is useful for plotting -fitts output from 3dDeconvolve and visually examining time series fitting
- Compute HRFs and statistics: 3dDeconvolve

### Script - 3dToutcount

```
# set list of runs
set runs = (`count -digits 2 1 10`)
# run 3dToutcount for each run
foreach run ( $runs )
  3dToutcount -automask pb00.$subj.r$run.tcat+orig > outcount r$run.1D
end
        10
             20
                  30.
                       40
                           50
                                          80
                                                    100.
                                                        110.
                                                             120.
                        Via 1dplot outcount r??.1D
```

3dToutcount searches for "outliers" in data time series; You should examine noticeable runs & time points

## Script - 3dTshift

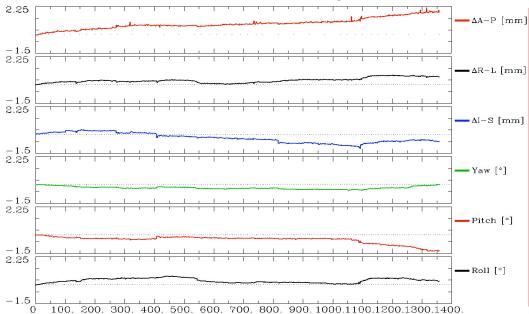
```
# run 3dTshift for each run
foreach run ( $runs )
   3dTshift -tzero 0 -quintic -prefix pb01.$subj.r$run.tshift \
        pb00.$subj.r$run.tcat+orig
end
```

- Produces new datasets where each time series has been shifted to have the same time origin
- -tzero 0 means that all data time series are interpolated to match the time offset of the first slice
  - Which is what the slice timing files usually refer to
  - Quintic (5<sup>th</sup> order) polynomial interpolation is used
- 3dDeconvolve will be run on these time-shifted datasets
  - This is mostly important for Event-Related FMRI studies, where the response to the stimulus is briefer than for Block designs
    - (Because the stimulus is briefer)
  - Being a little off in the stimulus timing in a Block design is not likely to matter much

## Script - 3dvolreg

#### end

- Produces new datasets where each volume (one time point) has been aligned (registered) to the #0 time point in the #1 dataset
- Movement parameters are saved into files dfile.r\$run.1D
  - Will be used as extra regressors in 3dDeconvolve to reduce motion artifacts



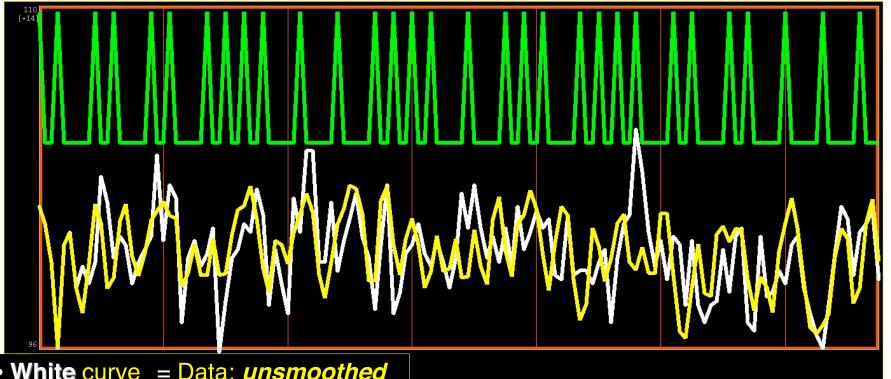
#### 1dplot -volreg dfile.rall.1D

- Shows movement parameters for all runs (1360 time points) in degrees and millimeters
- Very important to look at this graph!
- Excessive movement can make an imaging run useless — 3dvolreg won't be able to compensate
  - Pay attention to scale of movements: more than about 2 voxel sizes in a short time interval is usually bad

## Script - 3dmerge

```
# blur each volume
foreach run ( $runs )
    3dmerge -1blur fwhm 4 -doall -prefix pb03.$subj.r$run.blur
            pb02.$subj.r$run.volreg+orig
end
```

• Why Blur? Reduce noise by averaging neighboring voxels time series

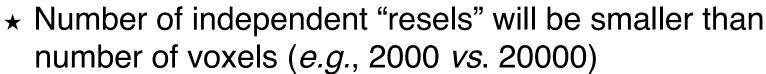


- White curve = Data: unsmoothed
- **llow** curve = Model fit  $(R^2 = 0.50)$   $\leftarrow$
- Green curve = Stimulus timing

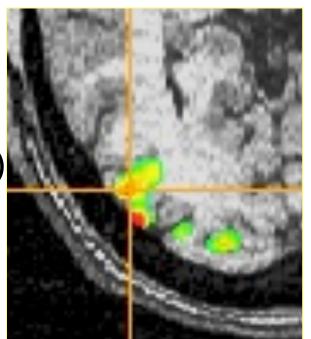
This is an extremely good fit for ER FMRI data!

### **Why Blur? - 2**

- fMRI activations are (usually)
   blob-ish (several voxels across)
- Averaging neighbors will also reduce the fiendish multiple comparisons problem



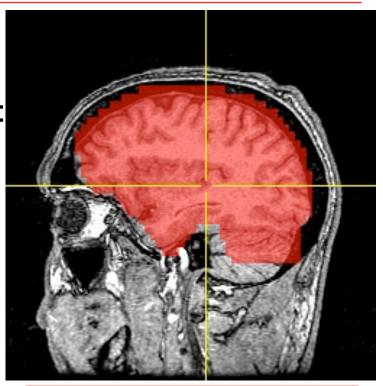
- Why not just acquire at lower resolution?
  - ⋆ To avoid averaging across brain/non-brain interfaces
  - ⋆ To project onto surface models
- Amount to blur is specified as FWHM (Full Width at Half Maximum) of spatial averaging filter (4 mm in script)



### Script - 3dAutomask

```
# create 'full_mask' dataset (union mask)
foreach run ( $runs )
    3dAutomask -dilate 1 -prefix rm.mask_r$run pb03.$subj.r$run.blur+orig
end
# get mean and compare it to 0 for taking 'union'
3dMean -datum short -prefix rm.mean rm.mask*.HEAD
3dcalc -a rm.mean+orig -expr 'ispositive(a-0)' -prefix full mask.$subj
```

- 3dAutomask creates a mask of contiguous high-intensity voxels (with some hole-filling) for each run separately: common mask for all runs/subjects
- 3dMean and 3dcalc are used to create a mask that is the <u>union</u> of all the individual run masks
- 3dDeconvolve analysis will be limited to voxels in this mask
  - Will run faster, since less data to process



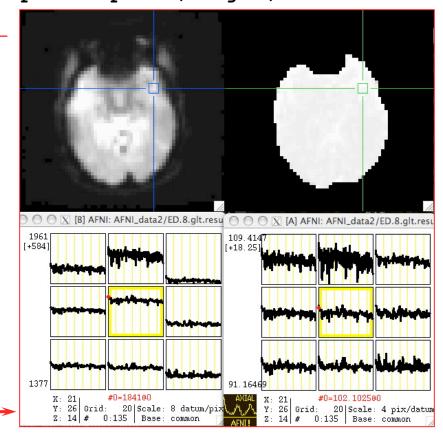
Automask from EPI shown in red

## Script - Scaling

```
# scale each voxel time series to have a mean of 100
# (subject to maximum value of 200)
foreach run ( $runs )
   3dTstat -prefix rm.mean_r$run pb03.$subj.r$run.blur+orig
   3dcalc -a pb03.$subj.r$run.blur+orig -b rm.mean_r$run+orig \
        -c full_mask.$subj+orig \
        -expr 'c * min(200, a/b*100)' -prefix pb04.$subj.r$run.scale
```

#### end

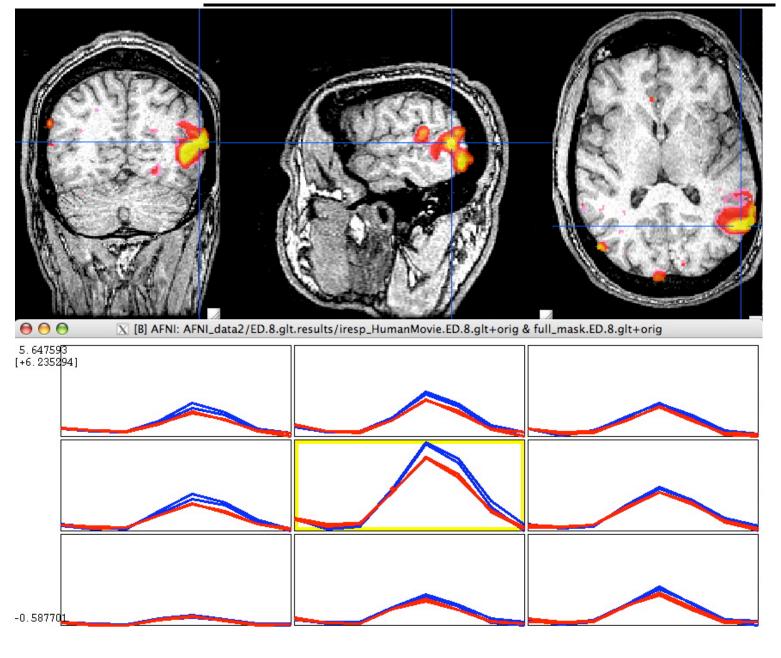
- 3dTstat calculates the mean (through time) of each voxel's time series data
- For voxels in the mask, each data point is scaled (multiplied) using 3dcalc so that it's time series will have mean=100
- If an HRF regressor has max amplitude = 1, then its  $\beta$  coefficient will represent the % signal change (from the mean) due to that part of the signal model
- Scaled images are very boring to view
  - No spatial contrast by design!
  - Graphs have common baseline now –



#### Script - 3dDeconvolve

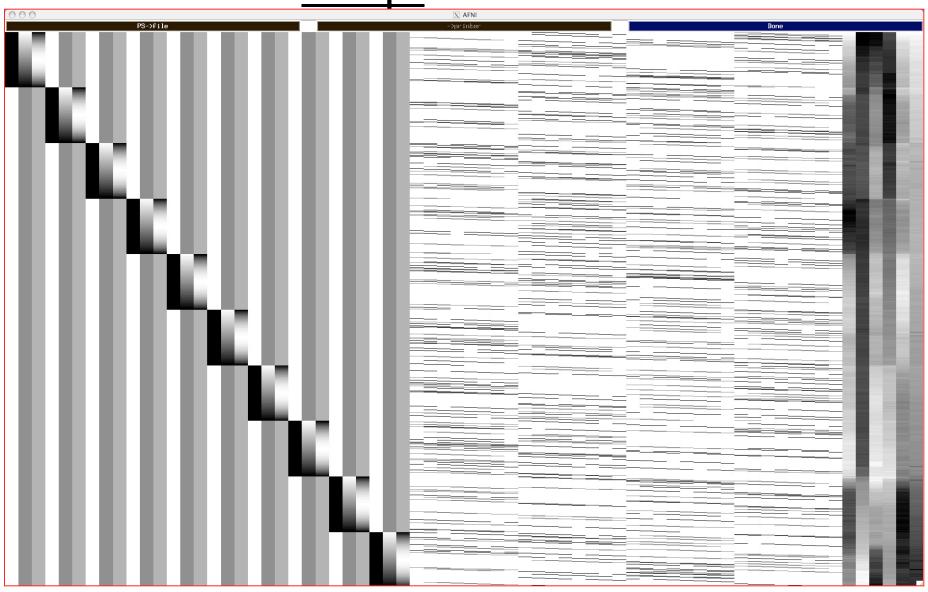
```
3dDeconvolve -input pb04.$subj.r??.scale+orig.HEAD -polort 2
 -mask full mask.$subj+orig -basis normall 1 -num stimts 10
 -stim times 1 stimuli/stim times.01.1D 'TENT(0,14,8)'
 -stim label 1 ToolMovie
 -stim times 2 stimuli/stim times.02.1D 'TENT(0,14,8)'
 -stim label 2 HumanMovie
                                                                        4 stim types
 -stim times 3 stimuli/stim times.03.1D 'TENT(0,14,8)'
 -stim label 3 ToolPoint
 -stim times 4 stimuli/stim times.04.1D 'TENT(0,14,8)'
 -stim label 4 HumanPoint
 -stim file 5 dfile.rall.1D'[0]' -stim base 5 -stim label 5 roll
 -stim file 6 dfile.rall.1D'[1]' -stim base 6 -stim label 6 pitch \
 -stim file 7 dfile.rall.1D'[2]' -stim base 7 -stim label 7 yaw
                                                                        motion params
 -stim file 8 dfile.rall.1D'[3]' -stim base 8 -stim label 8 dS
 -stim file 9 dfile.rall.1D'[4]' -stim base 9 -stim label 9 dL
 -stim file 10 dfile.rall.1D'[5]' -stim base 10 -stim label 10 dP
 -iresp 1 iresp ToolMovie.$subj -iresp 2 iresp HumanMovie.$subj
                                                                        HRF outputs
 -iresp 3 iresp ToolPoint.$subj -iresp 4 iresp HumanPoint.$subj
 -gltsym ../misc files/glt1.txt -glt label 1 FullF
 -gltsym ../misc files/glt2.txt -glt label 2 HvsT
 -gltsym ../misc files/glt3.txt -glt label 3 MvsP
                                                                        GLTs
 -qltsym ../misc files/qlt4.txt -qlt label 4 HMvsHP
 -gltsym ../misc files/glt5.txt -glt label 5 TMvsTP
 -gltsym ../misc files/glt6.txt -glt label 6 HPvsTP
 -gltsym ../misc files/glt7.txt -glt label 7 HMvsTM
 -fout -tout -full first -x1D Xmat.x1D -fitts fitts.$subj -bucket stats.$subj
```

#### Results: Humans vs. Tools



- Color overlay:HvsTGLTcontrast
- Blue (upper) graphs: Human HRFs
- Red (lower) graphs: Tool HRFs

# Script - X Matrix



Via 1grayplot -sep Xmat.x1D

#### Script - Random Comments

- •-polort 2
  - ★Sets baseline (detrending) to use quadratic polynomials—in each run
- •-mask full\_mask.\$subj+orig
  - ★Process only the voxels that are nonzero in this mask dataset
- -basis normall 1
  - ★Make sure that the basis functions used in the HRF expansion all have maximum magnitude=1
- -stim\_times 1 stimuli/stim\_times.01.1D
  'TENT(0,14,8)'
  - -stim label 1 ToolMovie
    - ★The HRF model for the ToolMovie stimuli starts at 0 s after each stimulus, lasts for 14 s, and has 8 (6 full + 2 half) basis tents
      ₀ Which have knots (breakpoints) spaced 14/(8-1) = 2 s apart
- •-iresp 1 iresp\_ToolMovie.\$subj
  - **★**The HRF model for the **ToolMovie** stimuli is output into dataset iresp\_**ToolMovie**.**ED**.8.glt+orig

## Script - GLTs

- -gltsym ../misc\_files/glt2.txt -glt\_label 2 HvsT
  - File ../misc files/glt2.txt contains 1 line of text:
    - o -ToolMovie +HumanMovie -ToolPoint +HumanPoint
    - This is the "Humans vs. Tools" HvsT contrast shown on Results slide
- This GLT means to take all 8  $\beta$  coefficients for each stimulus class and combine them with additions and subtractions as ordered:

$$LC = -\beta_0^{TM} - \dots - \beta_7^{TM} + \beta_0^{HM} + \dots + \beta_7^{HM} - \beta_0^{TP} - \dots - \beta_7^{TP} + \beta_0^{HP} + \dots + \beta_7^{HP}$$

- This test is looking at integrated (summed) response to "Human" stimuli and subtracting it from integrated response to "Tool" stimuli
- Combining subsets of the 
   β weights is also possible with -gltsym:
  - +HumanMovie[2..6] -HumanPoint[2..6]
  - This GLT would add up #2,3,4,5, & 6  $\beta$ s for one type of stimulus and subtract sum of #2,3,4,5, & 6  $\beta$ s for another type of stimulus
    - And also produce F- and t-statistics for this linear combination
  - Alternatively F-test for composite hypothesis: -gltsym `+HumanMovie [[2..6]] -HumanPoint[[2..6]]' -glt\_label 2 HvsT
     Θ<sub>2</sub><sup>TM</sup> = β<sub>2</sub><sup>TP</sup>, β<sub>3</sub><sup>TM</sup> = β<sub>3</sub><sup>TP</sup>, ..., β<sub>6</sub><sup>TM</sup> = β<sub>6</sub><sup>TP</sup>

## Script - Multi-Row GLTs

- GLTs presented up to now have had one row
  - $\star$  Testing if some linear combination of  $\beta$  weights is nonzero; test statistic is t or  $F(F=t^2)$  when testing a single number
  - ★ Testing if the X matrix columns, when added together to form one column as specified by the GLT (+ and -), explain a significant fraction of the data time series (equivalent to above)
- Can also do a single test to see if several different combinations of **b** weights are **all** zero +ToolMovie +HumanMovie -gltsym ../misc files/glt1.txt

```
-glt label 1 FullF
```

★ Tests if any of the stimulus classes have nonzero integrated HRF (each name means "add up those  $\beta$  weights"): DOF = (4,1292)

**+ToolPoint** 

+HumanPoint

★ Different than the default "Full F-stat" produced by 3dDeconvolve, which tests if any of the *individual*  $\beta$ weights are nonzero: DOF = (32, 1292)

#### Two Possible Formats for -stim times

19.4

- If you don't use -local\_times or -global\_times,
   3dDeconvolve will guess which way to interpret numbers:
- A single column of numbers (GLOBAL times)
  - ⋆ One stimulus time per row
  - $\star$  Times are relative to first image in dataset being at t=0
  - ★ May not be simplest to use if multiple runs are catenated
- One row for each run within a catenated dataset (LOCAL times)
  - ★ Each time in  $j^{th}$  row is relative to start of run #j being t=0
  - ★ If some run has NO stimuli in the given class, just put a single "\*" in that row as a filler
    4.7 9.6 11.8 19.4
    - Different numbers of stimuli per run are OK
    - At least one row must have more than 1 time
       (so that the LOCAL type of timing file can be told from the GLOBAL)
- Two methods are available because of users' diverse needs
  - ★ N.B.: if you chop first few images off the start of each run, the inputs to -stim\_times must be adjusted accordingly!
    - o Better to use -CENSORTR to tell 3dDeconvolve just to ignore those points

More information about-stim times and its variants is in the afni07 advanced talk

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